

# Supplementary Material

## Associations Between Caffeine Consumption, Cognitive Decline, and Dementia: A Systematic Review

**Supplementary Table 1.** The PRISMA checklist

Section/topic	#	Checklist item	Reported on page #
<b>TITLE</b>			
Title	1	Identify the report as a systematic review, meta-analysis, or both.	1
<b>ABSTRACT</b>			
Structured summary	2	Provide a structured summary including, as applicable: background; objectives; data sources; study eligibility criteria, participants, and interventions; study appraisal and synthesis methods; results; limitations; conclusions and implications of key findings; systematic review registration number.	1
<b>INTRODUCTION</b>			
Rationale	3	Describe the rationale for the review in the context of what is already known.	3
Objectives	4	Provide an explicit statement of questions being addressed with reference to participants, interventions, comparisons, outcomes, and study design (PICOS).	4
<b>METHODS</b>			
Protocol and registration	5	Indicate if a review protocol exists, if and where it can be accessed (e.g., Web address), and, if available, provide registration information including registration number.	NA
Eligibility criteria	6	Specify study characteristics (e.g., PICOS, length of follow-up) and report characteristics (e.g., years considered, language, publication status) used as criteria for eligibility, giving rationale.	Table 1
Information sources	7	Describe all information sources (e.g., databases with dates of coverage, contact with study authors to identify additional studies) in the search and date last searched.	4
Search	8	Present full electronic search strategy for at least one database, including any limits used, such that it could be repeated.	4
Study selection	9	State the process for selecting studies (i.e., screening, eligibility, included in systematic review, and, if applicable, included in the meta-analysis).	4
Data collection process	10	Describe method of data extraction from reports (e.g., piloted forms, independently, in duplicate) and any processes for obtaining and confirming data from investigators.	4

Data items	11	List and define all variables for which data were sought (e.g., PICOS, funding sources) and any assumptions and simplifications made.	4
Risk of bias in individual studies	12	Describe methods used for assessing risk of bias of individual studies (including specification of whether this was done at the study or outcome level), and how this information is to be used in any data synthesis.	5
Summary measures	13	State the principal summary measures (e.g., risk ratio, difference in means).	6
Synthesis of results	14	Describe the methods of handling data and combining results of studies, if done, including measures of consistency (e.g., $I^2$ ) for each meta-analysis.	6
Risk of bias across studies	15	Specify any assessment of risk of bias that may affect the cumulative evidence (e.g., publication bias, selective reporting within studies).	5
Additional analyses	16	Describe methods of additional analyses (e.g., sensitivity or subgroup analyses, meta-regression), if done, indicating which were pre-specified.	NA
<b>RESULTS</b>			
Study selection	17	Give numbers of studies screened, assessed for eligibility, and included in the review, with reasons for exclusions at each stage, ideally with a flow diagram.	7
Study characteristics	18	For each study, present characteristics for which data were extracted (e.g., study size, PICOS, follow-up period) and provide the citations.	Table 1
Risk of bias within studies	19	Present data on risk of bias of each study and, if available, any outcome level assessment (see item 12).	Supplementary Table 3
Results of individual studies	20	For all outcomes considered (benefits or harms), present, for each study: (a) simple summary data for each intervention group (b) effect estimates and confidence intervals, ideally with a forest plot.	Table 1
Synthesis of results	21	Present results of each meta-analysis done, including confidence intervals and measures of consistency.	NA
Risk of bias across studies	22	Present results of any assessment of risk of bias across studies (see Item 15).	8
Additional analysis	23	Give results of additional analyses, if done (e.g., sensitivity or subgroup analyses, meta-regression [see Item 16]).	NA
<b>DISCUSSION</b>			
Summary of evidence	24	Summarize the main findings including the strength of evidence for each main outcome; consider their relevance to key groups (e.g., healthcare providers, users, and policy makers).	14
Limitations	25	Discuss limitations at study and outcome level (e.g., risk of bias), and at review-level (e.g., incomplete retrieval of identified research, reporting bias).	16
Conclusions	26	Provide a general interpretation of the results in the context of other evidence, and implications for future research.	17
<b>FUNDING</b>			
Funding	27	Describe sources of funding for the systematic review and other support (e.g., supply of data); role of funders for the systematic review.	NA

**Supplementary Table 2.** Studies excluded on full-text level (n=44)

References	Reason for exclusion
1. An, R., Liu, G., Khan, N., Yan, H., & Wang, Y. (2019). Dietary Habits and Cognitive Impairment Risk Among Oldest-Old Chinese. <i>Journals of Gerontology Series B-Psychological Sciences and Social Sciences</i> , 74(3), 474–483. <a href="https://doi.org/10.1093/geronb/gbw170">https://doi.org/10.1093/geronb/gbw170</a>	Sharing population with another study
2. Bawa, S. (2009). The significance of polyphenols in the prevention of neurodegenerative diseases. <i>Agro Food Industry Hi-Tech</i> , 20(1), 42–46.	Study design
3. Borgwardt, S., Hammann, F., Scheffler, K., Kreuter, M., Drewe, J., & Beglinger, C. (2012). Neural effects of green tea extract on dorsolateral prefrontal cortex. <i>European Journal of Clinical Nutrition</i> , 66(11), 1187–1192. <a href="https://doi.org/10.1038/ejcn.2012.105">https://doi.org/10.1038/ejcn.2012.105</a>	No suitable cognitive outcome measure
4. Caparros-Lefebvre, D., Elbaz, A., & Grp, C. P. S. (1999). Possible relation of atypical parkinsonism in the French West Indies with consumption of tropical plants: a case-control study. <i>Lancet</i> , 354(9175), 281–286. <a href="https://doi.org/10.1016/S0140-6736(98)10166-6">https://doi.org/10.1016/S0140-6736(98)10166-6</a>	Non-caffeine effect
5. Carmichael, O. T., Pillai, S., Shankapal, P., McLellan, A., Kay, D. G., Gold, B. T., & Keller, J. N. (2018). A Combination of Essential Fatty Acids, Panax Ginseng Extract, and Green Tea Catechins Modifies Brain fMRI Signals in Healthy Older Adults. <i>The Journal of Nutrition, Health &amp; Aging</i> , 22(7), 837–846. <a href="https://doi.org/10.1007/s12603-018-1028-2">https://doi.org/10.1007/s12603-018-1028-2</a>	Combined intervention
6. Chin, A.-V., Robinson, D. J., O'Connell, H., Hamilton, F., Bruce, I., Coen, R., Walsh, B., Coakley, D., Molloy, A., Scott, J., Lawlor, B. A., & Cunningham, C. J. (2008). Vascular biomarkers of cognitive performance in a community-based elderly population: the Dublin Healthy Ageing study. <i>Age and Ageing</i> , 37(5), 559–564. <a href="https://doi.org/10.1093/ageing/afn144">https://doi.org/10.1093/ageing/afn144</a>	Non-caffeine effect
7. Commenges, D., Scotet, V., Renaud, S., Jacqmin-Gadda, H., Barberger-Gateau, P., & Dartigues, J. F. (2000). Intake of flavonoids and risk of dementia. <i>European Journal of Epidemiology</i> , 16(4), 357–363. <a href="https://doi.org/10.1023/A:1007614613771">https://doi.org/10.1023/A:1007614613771</a>	Combined intervention
8. Eskelinen, M. H., & Kivipelto, M. (2010). Caffeine as a protective factor in dementia and Alzheimer's disease. <i>Journal of Alzheimer's Disease</i> , 20 Suppl 1, S167-74. <a href="https://doi.org/10.3233/JAD-2010-1404">https://doi.org/10.3233/JAD-2010-1404</a>	Study design
9. Feng, L., Chong, M.-S., Lim, W.-S., Gao, Q., Nyunt, M. S. Z., Lee, T.-S., Collinson, S. L., Tsoi, T., Kua, E.-H., & Ng, T.-P. (2016). Tea consumption reduces the incidence of neurocognitive disorders: findings from the Singapore Longitudinal Aging Study. <i>Journal of Nutrition Health &amp; Aging</i> , 20(10), 1002–1009. <a href="https://doi.org/10.1007/s12603-016-0687-0">https://doi.org/10.1007/s12603-016-0687-0</a>	Sharing population with another study

10. Feng, L., Gwee, X., Kua, E.-H., & Ng, T.-P. (2010). Cognitive function and tea consumption in community dwelling older Chinese in Singapore. <i>Journal of Nutrition Health &amp; Aging</i> , 14(6), 433–438.	Sharing population with another study
11. Haller, S., Montandon, M.-L., Rodriguez, C., Moser, D., Toma, S., Hofmeister, J., & Giannakopoulos, P. (2017). Caffeine impact on working memory-related network activation patterns in early stages of cognitive decline. <i>Neuroradiology</i> , 59(4), 387–395. <a href="https://doi.org/10.1007/s00234-017-1803-5">https://doi.org/10.1007/s00234-017-1803-5</a>	No suitable cognitive outcome measure
12. Haller, S., Montandon, M.-L., Rodriguez, C., Moser, D., Toma, S., Hofmeister, J., Sinanaj, I., Lovblad, K.-O., & Giannakopoulos, P. (2014). Acute caffeine administration effect on brain activation patterns in mild cognitive impairment. <i>Journal of Alzheimer's Disease</i> , 41(1), 101–112. <a href="https://doi.org/10.3233/JAD-132360">https://doi.org/10.3233/JAD-132360</a>	No suitable cognitive outcome measure
13. Hosking, D. E., Nettelbeck, T., Wilson, C., & Danthiir, V. (2014). Retrospective lifetime dietary patterns predict cognitive performance in community-dwelling older Australians. <i>The British Journal of Nutrition</i> , 112(2), 228–237. <a href="https://doi.org/10.1017/S0007114514000646">https://doi.org/10.1017/S0007114514000646</a>	Combined intervention
14. Kim, J. W., Byun, M. S., Yi, D., Lee, J. H., Jeon, S. Y., Jung, G., Lee, H. N., Sohn, B. K., Lee, J.-Y., Kim, Y. K., Shin, S. A., Sohn, C.-H., & Lee, D. Y. (2019). Coffee intake and decreased amyloid pathology in human brain. <i>Translational Psychiatry</i> , 9(1), 270. <a href="https://doi.org/10.1038/s41398-019-0604-5">https://doi.org/10.1038/s41398-019-0604-5</a>	No suitable cognitive outcome measure
15. Kim, J., Yu, A., Choi, B. Y., Nam, J. H., Kim, M. K., Oh, D. H., & Yang, Y. J. (2015). Dietary Patterns Derived by Cluster Analysis are Associated with Cognitive Function among Korean Older Adults. <i>Nutrients</i> , 7(6), 4154–4169. <a href="https://doi.org/10.3390/nu7064154">https://doi.org/10.3390/nu7064154</a>	Combined intervention
16. Koppelstaetter, F., Poeppl, T. D., Siedentopf, C. M., Ischebeck, A., Kolbitsch, C., Mottaghay, F. M., Felber, S. R., Jaschke, W. R., & Krause, B. J. (2010). Caffeine and cognition in functional magnetic resonance imaging. <i>Journal of Alzheimer's Disease</i> , 20 Suppl 1, S71-84. <a href="https://doi.org/10.3233/JAD-2010-1417">https://doi.org/10.3233/JAD-2010-1417</a>	Study design
17. Kwok, M. K., Leung, G. M., & Schooling, C. M. (2016). Habitual coffee consumption and risk of type 2 diabetes, ischemic heart disease, depression and Alzheimer's disease: a Mendelian randomization study. <i>Scientific Reports</i> , 6, 36500. <a href="https://doi.org/10.1038/srep36500">https://doi.org/10.1038/srep36500</a>	Non-caffeine effect
18. Kyle, J., Fox, H. C., & Whalley, L. J. (2010). Caffeine, cognition, and socioeconomic status. <i>Journal of Alzheimer's Disease</i> , 20 Suppl 1, S151-9. <a href="https://doi.org/10.3233/JAD-2010-1409">https://doi.org/10.3233/JAD-2010-1409</a>	Sharing population with another study
19. Lang, R., Dieminger, N., Beusch, A., Lee, Y.-M., Dunkel, A., Suess, B., Skurk, T., Wahl, A., Hauner, H., & Hofmann, T. (2013). Bioappearance and pharmacokinetics of bioactives upon coffee consumption. <i>Analytical and Bioanalytical Chemistry</i> , 405(26), 8487–8503. <a href="https://doi.org/10.1007/s00216-013-7288-0">https://doi.org/10.1007/s00216-013-7288-0</a>	No suitable cognitive outcome measure
20. Laurin, D., Masaki, K. H., Foley, D. J., White, L. R., & Launer, L. J. (2004). Midlife dietary intake of antioxidants and risk of late-life incident dementia: the Honolulu-Asia Aging Study. <i>American</i>	Combined intervention

<p><i>Journal of Epidemiology</i>, 159(10), 959–967. <a href="https://doi.org/10.1093/aje/kwh124">https://doi.org/10.1093/aje/kwh124</a></p>	
21. Lefevre-Arbogast, S., Gaudout, D., Bensalem, J., Letenneur, L., Dartigues, J.-F., Hejblum, B. P., Feart, C., Delcourt, C., & Samieri, C. (2018). Pattern of polyphenol intake and the long-term risk of dementia in older persons. <i>Neurology</i> , 90(22), E1979–E1988. <a href="https://doi.org/10.1212/WNL.00000000000005607">https://doi.org/10.1212/WNL.00000000000005607</a>	Combined intervention
22. Low, D. Y., Lefèvre-Arbogast, S., González-Domínguez, R., Urpi-Sarda, M., Micheau, P., Petera, M., Centeno, D., Durand, S., Pujos-Guillot, E., Korosi, A., Lucassen, P. J., Aigner, L., Proust-Lima, C., Hejblum, B. P., Helmer, C., Andres-Lacueva, C., Thuret, S., Samieri, C., & Manach, C. (2019). Diet-Related Metabolites Associated with Cognitive Decline Revealed by Untargeted Metabolomics in a Prospective Cohort. <i>Molecular Nutrition &amp; Food Research</i> , 63(18), e1900177. <a href="https://doi.org/10.1002/mnfr.201900177">https://doi.org/10.1002/mnfr.201900177</a>	Non-caffeine effect
23. Moreira, A., Diógenes, M. J., de Mendonça, A., Lunet, N., & Barros, H. (2016). Chocolate Consumption is Associated with a Lower Risk of Cognitive Decline. <i>Journal of Alzheimer's Disease</i> , 53(1), 85–93. <a href="https://doi.org/10.3233/JAD-160142">https://doi.org/10.3233/JAD-160142</a>	Non-caffeine effect
24. Moss, M., Jones, R., Moss, L., Cutter, R., & Wesnes, K. (2016). Acute consumption of Peppermint and Chamomile teas produce contrasting effects on cognition and mood in healthy young adults. <i>Plant Science Today</i> , 3(3), 327–336. <a href="https://doi.org/10.14719/pst.2016.3.3.246">https://doi.org/10.14719/pst.2016.3.3.246</a>	Non-caffeine effect
25. Nutaitis, A. C., Tharwani, S. D., Serra, M. C., Goldstein, F. C., Zhao, L., Sher, S. S., Verble, D. D., & Wharton, W. (2019). Diet as a Risk Factor for Cognitive Decline in African Americans and Caucasians with a Parental History of Alzheimer's Disease: A Cross-Sectional Pilot Study Dietary Patterns. <i>The Journal of Prevention of Alzheimer's Disease</i> , 6(1), 50–55. <a href="https://doi.org/10.14283/jpad.2018.44">https://doi.org/10.14283/jpad.2018.44</a>	Non-caffeine effect
26. Okubo, H., Inagaki, H., Gondo, Y., Kamide, K., Ikebe, K., Masui, Y., Arai, Y., Ishizaki, T., Sasaki, S., Nakagawa, T., Kabayama, M., Sugimoto, K., Rakugi, H., & Maeda, Y. (2017). Association between dietary patterns and cognitive function among 70-year-old Japanese elderly: a cross-sectional analysis of the SONIC study. <i>Nutrition Journal</i> , 16(1), 56. <a href="https://doi.org/10.1186/s12937-017-0273-2">https://doi.org/10.1186/s12937-017-0273-2</a>	Combined intervention
27. Park, J., Han, J. W., Lee, J. R., Byun, S., Suh, S. W., Kim, J. H., & Kim, K. W. (2020). Association between lifetime coffee consumption and late life cerebral white matter hyperintensities in cognitively normal elderly individuals. <i>Scientific Reports</i> , 10(1), 421. <a href="https://doi.org/10.1038/s41598-019-57381-z">https://doi.org/10.1038/s41598-019-57381-z</a>	No suitable cognitive outcome measure
28. Park, S.-K., Jung, I.-C., Lee, W. K., Lee, Y. S., Park, H. K., Go, H. J., Kim, K., Lim, N. K., Hong, J. T., Ly, S. Y., & Rho, S. S. (2011). A Combination of Green Tea Extract and L-Theanine Improves Memory and Attention in Subjects with Mild Cognitive Impairment: A Double-Blind Placebo-Controlled Study. <i>Journal of</i>	Non-caffeine effect

<i>Medicinal Food</i> , 14(4), 334–343. <a href="https://doi.org/10.1089/jmf.2009.1374">https://doi.org/10.1089/jmf.2009.1374</a>	
29. Pin, N. T. (2016). Cognitive Health of Older Persons in Longitudinal Ageing Cohort Studies. <i>Sains Malaysiana</i> , 45(9), 1351–1355.	Study design
30. Ritchie, K., Ancelin, M. L., Amieva, H., Rouaud, O., & Carriere, I. (2014). The association between caffeine and cognitive decline: examining alternative causal hypotheses. <i>International Psychogeriatrics</i> , 26(4), 581–590. <a href="https://doi.org/10.1017/S1041610213002469">https://doi.org/10.1017/S1041610213002469</a>	No suitable cognitive outcome measure
31. Ritchie, K., Artero, S., Portet, F., Brickman, A., Muraskin, J., Beanino, E., Ancelin, M.-L., & Carrière, I. (2010). Caffeine, cognitive functioning, and white matter lesions in the elderly: establishing causality from epidemiological evidence. <i>Journal of Alzheimer's Disease</i> , 20 Suppl 1, S161-6. <a href="https://doi.org/10.3233/JAD-2010-1387">https://doi.org/10.3233/JAD-2010-1387</a>	Sharing population with another study
32. Robinson, J. L., Hunter, J. M., Reyes-Izquierdo, T., Argumedo, R., Brizuela-Bastien, J., Keller, R., & Pietrzkowski, Z. J. (n.d.). Cognitive short- and long-term effects of coffee cherry extract in older adults with mild cognitive decline. <i>Aging Neuropsychology and Cognition</i> . <a href="https://doi.org/10.1080/13825585.2019.1702622">https://doi.org/10.1080/13825585.2019.1702622</a>	Non-caffeine effect
33. Rodriguez-Bailon, M., Garcia-Moran, T., Montoro-Membila, N., Rodenas-Garcia, E., Arnedo Montoro, M., & Funes Molina, M. J. (2017). Positive and Negative Consequences of Making Coffee among Breakfast Related Irrelevant Objects: Evidence from MCI, Dementia, and Healthy Ageing. <i>Journal of the International Neuropsychological Society</i> , 23(6), 481–492. <a href="https://doi.org/10.1017/S135561771700025X">https://doi.org/10.1017/S135561771700025X</a>	Non-caffeine effect
34. Rubio-Perez, J. M., Albaladejo, M. D., Zafrilla, P., Vidal-Guevara, M. L., & Morillas-Ruiz, J. M. (2016). Effects of an antioxidant beverage on biomarkers of oxidative stress in Alzheimer's patients. <i>European Journal of Nutrition</i> , 55(6), 2105–2116. <a href="https://doi.org/10.1007/s00394-015-1024-9">https://doi.org/10.1007/s00394-015-1024-9</a>	No suitable cognitive outcome measure
35. Schmidt, A., Hammann, F., Wölnerhanssen, B., Meyer-Gerspach, A. C., Drewe, J., Beglinger, C., & Borgwardt, S. (2014). Green tea extract enhances parieto-frontal connectivity during working memory processing. <i>Psychopharmacology</i> , 231(19), 3879–3888. <a href="https://doi.org/10.1007/s00213-014-3526-1">https://doi.org/10.1007/s00213-014-3526-1</a>	No suitable cognitive outcome measure
36. Shirai, Y., Kuriki, K., Otsuka, R., Kato, Y., Nishita, Y., Tange, C., Tomida, M., Imai, T., Ando, F., & Shimokata, H. (2019). Association between green tea intake and risk of cognitive decline, considering glycated hemoglobin level, in older Japanese adults: the NILS-LSA study. <i>Nagoya Journal of Medical Science</i> , 81(4), 655–666. <a href="https://doi.org/10.18999/nagjms.81.4.655">https://doi.org/10.18999/nagjms.81.4.655</a>	Combined intervention
37. Su, N., Li, W., Li, X., Wang, T., Zhu, M., Liu, Y., Shi, Y., & Xiao, S. (2017). The Relationship between the Lifestyle of the Elderly in Shanghai Communities and Mild Cognitive Impairment. <i>Shanghai Archives of Psychiatry</i> , 29(6), 352–357. <a href="https://doi.org/10.11919/j.issn.1002-0829.217059">https://doi.org/10.11919/j.issn.1002-0829.217059</a>	Sharing population with another study

<p>38. Tomata, Y., Kakizaki, M., Nakaya, N., Tsuboya, T., Sone, T., Kuriyama, S., Hozawa, A., &amp; Tsuji, I. (2012). Green tea consumption and the risk of incident functional disability in elderly Japanese: the Ohsaki Cohort 2006 Study. <i>American Journal of Clinical Nutrition</i>, 95(3), 732–739.  <a href="https://doi.org/10.3945/ajcn.111.023200">https://doi.org/10.3945/ajcn.111.023200</a></p>	<p>No suitable cognitive outcome measure</p>
<p>39. Travassos, M., Santana, I., Baldeiras, I., Tsolaki, M., Gkatzima, O., Sermin, G., Yener, G. G., Simonsen, A., Hasselbalch, S. G., Kapaki, E., Mara, B., Cunha, R. A., Agostinho, P., Blennow, K., Zetterberg, H., Mendes, V. M., Manadas, B., &amp; de Mendon, A. (2015). Does Caffeine Consumption Modify Cerebrospinal Fluid Amyloid-<math>\beta</math> Levels in Patients with Alzheimer's Disease? <i>Journal of Alzheimer's Disease: JAD</i>, 47(4), 1069–1078.  <a href="https://doi.org/10.3233/JAD-150374">https://doi.org/10.3233/JAD-150374</a></p>	<p>No suitable cognitive outcome measure</p>
<p>40. Walters, E. R., &amp; Lesk, V. E. (2015). Time of day and caffeine influence some neuropsychological tests in the elderly. <i>Psychological Assessment</i>, 27(1), 161–168.  <a href="https://doi.org/10.1037/a0038213">https://doi.org/10.1037/a0038213</a></p>	<p>Sharing population with another study</p>
<p>41. Wang, H., Sun, W., Chang, Y., Wu, Z., Xu, Y., Wang, E., Wang, L., &amp; Yi, P. (2019). Effect of green tea consumption on human brain function in resting-state functional MRI. <i>Asia Pacific Journal of Clinical Nutrition</i>, 28(4), 740–746.  <a href="https://doi.org/10.6133/apjcn.201912_28(4).0010">https://doi.org/10.6133/apjcn.201912_28(4).0010</a></p>	<p>No suitable cognitive outcome measure</p>
<p>42. Wesnes, K. A., Barrett, M. L., &amp; Udani, J. K. (2013). An evaluation of the cognitive and mood effects of an energy shot over a 6 h period in volunteers. A randomized, double-blind, placebo controlled, cross-over study. <i>Appetite</i>, 67, 105–113.  <a href="https://doi.org/10.1016/j.appet.2013.04.005">https://doi.org/10.1016/j.appet.2013.04.005</a></p>	<p>Combined intervention</p>
<p>43. West, R. K., Ravona-Springer, R., Livny, A., Heymann, A., Shahar, D., Leroith, D., Preiss, R., Zukran, R., Silverman, J. M., &amp; Schnaider-Beeri, M. (2019). Age Modulates the Association of Caffeine Intake With Cognition and With Gray Matter in Elderly Diabetics. <i>The Journals of Gerontology. Series A, Biological Sciences and Medical Sciences</i>, 74(5), 683–688.  <a href="https://doi.org/10.1093/gerona/gly090">https://doi.org/10.1093/gerona/gly090</a></p>	<p>Combined intervention</p>
<p>44. Zhao, J., Zhang, X., &amp; Li, Z. (2019). The Relationship between Cognitive Impairment and Social Vulnerability among the Elderly: Evidence from an Unconditional Quantile Regression Analysis in China. <i>International Journal of Environmental Research and Public Health</i>, 16(19). <a href="https://doi.org/10.3390/ijerph16193684">https://doi.org/10.3390/ijerph16193684</a></p>	<p>Non-caffeine effect</p>

**Supplementary Table 3.** Risk of bias assessment for specific domains of ROBINS-I tool

Study	Bias							
	Confounding	Selection	Classification	Deviations	Missing	Measurements	Reporting	Overall
Assessment	Corrected Measured correctly Switching groups	Patient characteristics Start of intervention = start of study Correction for selection bias	Intervention defined Defined at start Affected by knowledge of outcome	Deviations intended intervention Unbalanced groups Co-interventions balanced Completed Adherence	Outcome available Participants excluded Balanced between groups Results robust to missing data	Influenced by intervention received Assessors aware of intervention Comparable across intervention groups Systematic errors	Selective reporting of outcomes Multiple analyses Subgroups	
<b>Al-khateeb et al. 2014</b>	PY	PN	PN	NI	PN	PY	PN	Moderate
<b>Arab et al. 2011</b>	N	N	PN	PY	PN	PN	PN	Low
<b>Arab et al. 2016</b>	Y	PY	PN	PY	PN	PY	PN	Serious
<b>Araújo et al. 2015</b>	N	PY	PN	PY	N	PN	PN	Low
<b>Araújo et al. 2016</b>	N	PN	N	PY	N	PY	N	Low
<b>Beydoun et al. 2014</b>	N	N	PN	PN	PN	PN	PN	Low
<b>Boot et al., 2013</b>	PN	N	PN	PY	PN	PN	PN	Low
<b>Broe et al. 1990</b>	PY	PY	N	PY	PN	PN	N	Moderate
<b>Cao et al. 2012</b>	Y	PN	PN	PN	PN	PN	PN	Moderate
<b>Chen et al. 2012</b>	N	PN	PN	PY	PN	PN	PN	Low
<b>Chin et al. 2008</b>	N	PN	PN	Y	PY	PN	PN	Moderate
<b>Cho et al. 2018</b>	PN	PY	PY	NA	PN	PN	PN	Moderate
<b>Chuang et al. 2019</b>	N	N	PN	PY	PN	PN	PY	Low
<b>Corley et al. 2010</b>	N	N	PN	PN	PY	PN	PN	Low
<b>Dai et al. 2006</b>	PN	PY	PN	PY	PN	PN	PN	Moderate
<b>Dong et al. 2020</b>	N	N	PN	PY	PN	PN	PN	Low
<b>Driscoll et al. 2016</b>	N	PY	PN	PY	PN	PY	PN	Moderate
<b>Eskelinen et al. 2009</b>	N	N	PN	PY	N	PY	N	Low
<b>Feng et al. 2012</b>	N	PY	PN	PY	PN	PY	PN	Moderate
<b>Feng et al. 2018</b>	PN	PN	PN	PN	N	PN	N	Low
<b>Fischer et al. 2018</b>	PN	PN	PY	PY	N	PN	PN	Moderate
<b>Gelber et al. 2011</b>	N	PN	PN	PY	PN	PN	N	Low
<b>Gu et al. 2018</b>	N	PN	PN	PY	PN	PN	PN	Low

Haller et al. 2018	N	PY	N	PY	PY	PN	PN	Moderate
Huang et al. 2009	N	PY	PN	PY	PN	PY	PY	Moderate
Ide et al. 2014	Y	PN	PN	PN	PY	PY	N	Moderate
Ide et al. 2016	N	PN	N	PY	PY	N	N	Low
Iranpour et al. 2019	PN	N	PN	PY	PN	PN	PN	Low
Jarvis, 1993	PN	N	PN	PY	PY	PN	PN	Moderate
Johnson-Kozlow et al. 2002	N	PN	PN	PY	PY	PN	PY	Moderate
Kitamura et al. 2016	N	PN	PN	PY	PY	PN	PY	Moderate
Konishi et al. 2018	PY	N	N	PN	PN	PN	PN	Low
Kuriyama et al. 2006	N	PN	PN	PY	PN	PN	PN	Low
Laitala et al. 2009	PN	N	N	PY	PN	PN	PY	Low
Lammi et al. 1989	Y	PY	PN	PY	PN	PN	PY	Serious
Larsson & Wolk, 2018	N	PN	N	PY	N	PN	N	Low
Lee et al. 2017	N	N	N	PY	PN	PN	PN	Low
Lesk et al. 2009	N	PN	PN	N	PN	PN	PN	Low
Lindsay et al. 2002	PN	PN	PN	PN	PN	PN	N	Low
Maia & de Mendonça, 2002	Y	PN	PN	PY	PN	N	N	Moderate
Mirza et al. 2014	N	PN	PN	PY	PN	N	PN	Low
Ng et al. 2008	N	PN	PN	PY	PN	PN	PN	Low
Noguchi-Shinohara et al. 2014	N	PN	PN	PY	PN	PN	PN	Low
Nurk et al. 2009	N	PN	PN	PY	PN	PN	PN	Low
Paganini-Hill et al. 2016	PN	PY	N	PY	N	PN	PN	Low
Richie et al., 2007	N	PN	PN	PN	PN	PN	PN	Low
Santos et al. 2010	N	PN	PN	PY	PN	PN	PY	Moderate
Shen et al. 2015	N	N	PN	PY	N	PN	N	Low
Shirai et al. 2020	N	N	PN	PY	PN	PY	PN	Low
Smith, 2009	PN	PN	PN	Y	PY	PN	PN	Moderate
Solfrizzi et al. 2015	N	N	PN	PY	PN	PN	PN	Low
Sugiyama et al. 2016	N	N	PN	PY	PN	PN	PN	Low
Tomata et al. 2016	N	N	PN	PY	PN	N	PN	Low
Tyas et al. 2001	N	PN	PN	PN	PN	PN	PN	Low
Valls-Pedret et al. 2012	PN	PN	PY	PY	PN	PN	PN	Moderate
van Boxtel et al. 2003	N	PN	PN	PY	PN	PN	N	Low
van Gelder et al. 2007	N	N	PN	PY	PN	PN	PN	Low
Vercambre et al. 2013	N	PY	PN	PY	PN	PN	PN	Moderate
Walters & Lesk, 2016	PN	N	PN	PY	PY	N	PN	Low
Wang et al. 2017	PY	PN	PN	Y	PN	PN	PY	Moderate

<b>Wu et al. 2011</b>	PN	N	PN	PN	PN	PN	N	Low
<b>Xu et al. 2018</b>	PN	PN	PY	PY	PN	PN	PY	Moderate
<b>Yang et al. 2016</b>	PN	PN	PN	PY	PN	PN	N	Low

Possible answers: NA (not applicable) NI (not assessable due to lack of info) N (no bias) PN (possibly no bias [uncertain]) PY (possibly bias [uncertain]) Y (bias)

5-point Likert scale: N -2 PN -1 NI/NA 0 PY 1 Y 2; lower (negative scores are better) and the higher the score, the more bias